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| APPLICATION NO. | FILING DATE | FIRST NAMED INVENTOR | ATTORNEY DOCKET NO. | CONFIRMATION NO. |
|-------------------------------------------------------------------------------------------------------|-------------|----------------------|---------------------|------------------|
| 10/788,793 | 02/27/2004 | Makoto Sato | 671302-2005 | 8148 |
| 7590 | 05/03/2005 | | EXAMINER | |
| THOMAS J. KOWALSKI, Esq. c/o FROMMER LAWRENCE & HAUG LLP 745 Fifth Avenue New York, NY 10151 | | | | ROBINSON, HOPE A |
| | | ART UNIT | | PAPER NUMBER |
| | | 1653 | | |

DATE MAILED: 05/03/2005

Please find below and/or attached an Office communication concerning this application or proceeding.

| | | |
|------------------------------|-------------------------------------|-------------------------|
| Office Action Summary | Application No. | Applicant(s) |
| | 10/788,793 | SATO ET AL. |
| | Examiner Hope A. Robinson | Art Unit 1653 |

-- The MAILING DATE of this communication appears on the cover sheet with the correspondence address --

Period for Reply

A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) FROM THE MAILING DATE OF THIS COMMUNICATION.

- Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication.
- If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely.
- If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication.
- Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133).
- Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).

Status

- 1) Responsive to communication(s) filed on 25 January 2005.
 2a) This action is FINAL. 2b) This action is non-final.
 3) Since this application is in condition for allowance except for formal matters, prosecution as to the merits is closed in accordance with the practice under *Ex parte Quayle*, 1935 C.D. 11, 453 O.G. 213.

Disposition of Claims

- 4) Claim(s) 1-27 is/are pending in the application.
 4a) Of the above claim(s) 4-12 and 14-27 is/are withdrawn from consideration.
 5) Claim(s) _____ is/are allowed.
 6) Claim(s) 1-3 and 13 is/are rejected.
 7) Claim(s) _____ is/are objected to.
 8) Claim(s) _____ are subject to restriction and/or election requirement.

Application Papers

- 9) The specification is objected to by the Examiner.
 10) The drawing(s) filed on 27 February 2004 is/are: a) accepted or b) objected to by the Examiner.
 Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).
 Replacement drawing sheet(s) including the correction is required if the drawing(s) is objected to. See 37 CFR 1.121(d).
 11) The oath or declaration is objected to by the Examiner. Note the attached Office Action or form PTO-152.

Priority under 35 U.S.C. § 119

- 12) Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).
 a) All b) Some * c) None of:
 1. Certified copies of the priority documents have been received.
 2. Certified copies of the priority documents have been received in Application No. _____.
 3. Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)).

* See the attached detailed Office action for a list of the certified copies not received.

Attachment(s)

- 1) Notice of References Cited (PTO-892)
 2) Notice of Draftsperson's Patent Drawing Review (PTO-948)
 3) Information Disclosure Statement(s) (PTO-1449 or PTO/SB/08)
 Paper No(s)/Mail Date 3/17/04.
- 4) Interview Summary (PTO-413)
 Paper No(s)/Mail Date, _____.
 5) Notice of Informal Patent Application (PTO-152)
 6) Other Alignments

DETAILED ACTION

Application Status

1. Applicant's election with traverse of Group I (claims 1-3 and 13) on January 25, 2005 is acknowledged. Claims 4-12 and 14-27 are withdrawn from further consideration pursuant to 37 CFR 1.12(b), as being drawn to a non-elected invention, there being no allowable generic or linking claim.

Restriction Requirement

2. The traversal is on the grounds that Groups I-VI should be rejoined because the groups are inextricably linked (i.e. Groups I and II); there is no serious burden of search (i.e Groups I-III); and that the search is coextensive (Groups IV-VI). Essentially Applicant argues that the relatedness between Groups means that a search of one would obtain the other, thus there is no burden of search. Applicant also argues that additional cost would be incurred to the applicant and patent office with the present restriction requirement. Cost is not germane to the issue of whether or not a restriction requirement is proper, therefore, no further comments will be made on this issue by the examiner. Regarding applicant's statement that there is no burden of search, the MPEP in chapter 800 indicates that the claimed invention by acquiring a separate status in the art demonstrates search burden. Furthermore, the search of the claimed invention is not coextensive as a reference that teaches one invention would not necessarily anticipate or make obvious another invention. However, if applicant is willing to make a

statement on the record that this is the case, it will be considered. The antibody, protein, DNA and non-human animal products are patentably distinct (Inventions I-V) as outlined in the previous office action, they have different structures, functions and modes of operation. For example, although the DNA encodes the protein, the DNA can be used to make probes or primers or used in a hybridization assay. Further, the protein can be used to make antibodies or in a bioassay. The method set forth in Invention VI is not the only process that the product can be used in. Moreover, MPEP chapter 800 state that a restriction requirement is proper if the inventions are independent or distinct (related or unrelated). Therefore the restriction requirement is deemed proper and is final.

Specification

3. The specification is objected to because of the following informalities:
 - (a) The specification is objected to because trademarks are disclosed throughout the instant specification and not all of them are capitalized or accompanied by the generic terminology. The use of the trademarks such as FIAGS™, GENBANK™, TRITON®-X-100, TRIS®, for example, have been noted in this application (see pages 17 and 32-34). It should be capitalized wherever it appears and be accompanied by the generic terminology. Although the use of trademarks is permissible in patent applications, the proprietary nature of the marks should be respected and every effort made to prevent their use in any manner, which might adversely affect their validity as trademarks.

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- (b) The specification is objected to because it contains an embedded hyperlink and/or other form of browser-executable code. Applicant is required to delete the embedded hyperlink and/or other form of browser-executable code. See MPEP § 608.01. See pages 14 (lines 13 and 25) and pages 15-17 for example. It is suggested that http:// is deleted.
- (c) The following typographical error appears on page 41 of the instant specification, "antiboies" which should be "antibodies".
- (d) The sequence notation throughout the instant specification is improper. See for example, "Seq. ID Nos. 5 and 6" on page 5 which should be "SEQ ID NOS:5 and 6".
- (e) The word "of" is missing following the word "consisting" on page 6, lines 11 and 22.
- (f) The specification is objected to because the claims appear on pages 42-46 of the instant specification instead of on a separate page. If applicant intends this to be text, the claim numbers should be removed.
- (g) The specification is objected to because "paragraphs" are referred to throughout the specification, see for example page 6 where the following appears "(paragraph 1); "(paragraph 2)" etc. Applicant is reminded of the proper format of the specification below.

The following guidelines illustrate the preferred layout for the specification of a utility application. These guidelines are suggested for the applicant's use.

Arrangement of the Specification

As provided in 37 CFR 1.77(b), the specification of a utility application should include the following sections in order. Each of the lettered items should appear in

upper case, without underlining or bold type, as a section heading. If no text follows the section heading, the phrase "Not Applicable" should follow the section heading:

- (a) TITLE OF THE INVENTION.
- (b) CROSS-REFERENCE TO RELATED APPLICATIONS.
- (c) STATEMENT REGARDING FEDERALLY SPONSORED RESEARCH OR DEVELOPMENT.
- (d) THE NAMES OF THE PARTIES TO A JOINT RESEARCH AGREEMENT
- (e) INCORPORATION-BY-REFERENCE OF MATERIAL SUBMITTED ON A COMPACT DISC (See 37 CFR 1.52(e)(5) and MPEP 608.05. Computer program listings (37 CFR 1.96(c)), "Sequence Listings" (37 CFR 1.821(c)), and tables having more than 50 pages of text are permitted to be submitted on compact discs.) or REFERENCE TO A "MICROFICHE APPENDIX" (See MPEP § 608.05(a). "Microfiche Appendices" were accepted by the Office until March 1, 2001.)
- (f) BACKGROUND OF THE INVENTION.
 - (1) Field of the Invention.
 - (2) Description of Related Art including information disclosed under 37 CFR 1.97 and 1.98.
- (g) BRIEF SUMMARY OF THE INVENTION.
- (h) BRIEF DESCRIPTION OF THE SEVERAL VIEWS OF THE DRAWING(S).
- (i) DETAILED DESCRIPTION OF THE INVENTION.
- (j) CLAIM OR CLAIMS (commencing on a separate sheet).
- (k) ABSTRACT OF THE DISCLOSURE (commencing on a separate sheet).
- (l) SEQUENCE LISTING (See MPEP § 2424 and 37 CFR 1.821-1.825. A "Sequence Listing" is required on paper if the application discloses a nucleotide or amino acid sequence as defined in 37 CFR 1.821(a) and if the required "Sequence Listing" is not submitted as an electronic document on compact disc).

Correction of the above and compliance with the sequence rules is required.

Drawing

4. It is noted that this application contains drawings executed in color (for example, Fig. 1), however, it does not appear that a petition to accept the drawings has been

filed. A decision on the petition will occur under separate cover once filed. Therefore, the drawings are objected to. Hence Figures such as Fig. 1 appears dark.

Information Disclosure Statement

5. The Information Disclosure Statement filed on March 17, 2004 has been received and entered. The references cited on the PTO-1449 Form have been considered by the examiner and a copy is attached to the instant Office action.

Claim Objection

6. Claims 3 and 13 is objected to because of the following informalities:

Claim 3 is objected to for the recitation of "65C" instead of "65°C".

Claim 13 is indefinite because the claim depends from a non-elected claim.

Correction of the above is required.

Claim Rejections - 35 USC § 112

The following is a quotation of the first paragraph of 35 U.S.C. 112:

The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.

7. Claims 1-3 and 13 are rejected under 35 U.S.C. 112, first paragraph, as containing subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventor(s), at the time the application was filed, had possession of the claimed invention.

The claimed invention is directed to an isolated DNA molecule that encodes a protein that controls cell migration and cell death wherein one or several amino acids are deleted, substituted or added or wherein all or part of the sequences is present or a DNA that hybridizes to the encoding DNA under stringent conditions and the conditions provided are merely exemplary, not limiting (see claims 1-3). The claims encompass a large genus that has not been adequately described. In addition, based on the open language "comprising", the claimed fragment is unlimited, thus having an undefined structure (see claim 1 for example). Note that claim 13 is directed to a host cell the comprise the encoded protein or a part of the protein and no description is provided as to what part of the protein the host cell is capable of expressing. Therefore, the claims read on several fragments, which have not been adequately described, and there is no indication as to a conserved region or where in the sequence the modifications could occur. Furthermore, the claims encompass a sequence that is completely deleted. Therefore, the skilled artisan cannot envision the detailed chemical structure of the claimed protein fragments, thus, claims reciting said protein fragments polypeptide lacks adequate written description. Additionally, claim 3 is directed to a DNA sequence that hybridizes under stringent conditions to the claimed DNA, however, the claim provides an example of stringent conditions, which is not limiting. It is known in the prior art that

hybridization conditions can vary; therefore, the claims need to recite the actual hybridization conditions. Further, the discussion provided in the instant specification does not breathe life into the claims. Note also that claim 2 recites the language "comprising part or all of either of these sequences" and the claim does not recite any functional language, thus said protein might not be biologically active or have a different activity other than cell migration and cell death control.

The instant specification disclose that the protein controls cell migration and cell death, however, there is no demonstration of a protein that has one or several amino acids deleted retaining this function. The specification lacks adequate written description for the claimed fragments thereof, with regard to size, structure and function (i.e. is function retained or is the fragment non-functional or possess a different function). The specification fails to provide any additional representative species of the claimed genus to show that applicant was in possession of the claimed genus. A representative number of species means that the species, which are adequately described, are representative of the entire genus. The written description requirement for a claimed genus may be satisfied through sufficient description of a representative number of species by actual reduction to practice, disclosure of drawings, or by disclosure of relevant identifying characteristics, for example, structure or other physical and/or chemical properties, by functional characteristics coupled with a known or disclosed correlation between function and structure, or by a combination of such identifying characteristics, sufficient to show the applicant was in possession of the claimed genus.

Accordingly, in the absence of sufficient recitation of distinguishing identifying characteristics, the specification does not provide adequate written description of the claimed genus. The claimed genus of protein fragments could include non-functional proteins or proteins with a different function than the one described. Therefore, the genus of claimed fragments encompasses widely variant species. As such, neither the description of the structure and function of SEQ ID NOS: 2, 4 and 6, for example, controlling cell migration and cell death is sufficient to be representative of the attributes and features of the entire genus. Based on the unlimited variations contemplated one skilled in the art would at best expect a protein that is different or at worst a protein that is not functional.

Vas-Cath Inc. v. Mahurkar, 935 F.2d 1555, 1563-64, 19 USPQ2d 1111, 1117 (Fed. Cir. 1991), states that "applicant must convey with reasonable clarity to those skilled in the art that, as of the filing date sought, he or she was in possession of the invention. The invention is, for purposes of the 'written description' inquiry, whatever is now claimed" (See page 1117). The specification does not "clearly allow persons of ordinary skill in the art to recognize that [he or she] invented what is claimed" (See *Vas-Cath* at page 1116). The skilled artisan cannot envision the detailed chemical structure of the encompassed genus of polypeptides, and therefore, conception is not achieved until reduction to practice has occurred, regardless of the complexity or simplicity of the method of isolation. Adequate written description requires more than a mere statement that it is part of the invention and reference to a potential method of isolating it. The

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compound itself is required. See *Fiers v. Revel*, 25 USPQ2d 1601 at 1606 (CAFC 1993).

Therefore, for all these reasons the specification lacks adequate written description, and one of skill in the art cannot reasonably conclude that the applicant had possession of the claimed invention at the time the instant application was filed.

8. Claims 1-3 and 13 are rejected under 35 U.S.C. 112, first paragraph, because the specification, while being enabling for the DNA encoding proteins set forth in SEQ ID NOS: 2, 4 and 6 that controls cell migration and cell death, does not reasonably provide enablement for any protein fragment thereof. The specification does not enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make the invention commensurate in scope with these claims. The enablement requirement refers to the requirement that the specification describe how to make and how to use the invention. There are many factors to be considered when determining whether there is sufficient evidence to support a determination that a disclosure does not satisfy the enablement requirement and whether any necessary experimentation is undue. These factors include, but are not limited to: Quantity of Experimentation Necessary; Amount of direction or guidance presented; Presence or absence of working examples; Nature of the Invention; State of the prior art and Relative skill of those in the art; Predictability or unpredictability of the art and Breadth of

the claims (see *In re Wands*, 858 F.2d at 737, 8 USPQ2d at 1404 (Fed. Cir. 1988). The factors most relevant to the instant invention are discussed below.

The amount of experimentation required to practice the claimed invention is undue as the claims encompass an unspecified amount of protein fragments, which may not retain the ascribed function. Based on the large amount of variability contemplated said protein fragment may not have the function ascribed to SEQ ID NOS: 2, 4 and 6 (controlling cell migration and cell death), however, the claims are directed to a DNA that encodes a protein comprising an amino acid sequence wherein 1 or several amino acids are deleted, substituted or added in SEQ ID NOS: 2, 4 and 6 and the fragments encompassed are not exemplified with the ascribed function. Thus the entire sequence could be substituted or deleted as there is limitation on the amount of residues that could be deleted or substituted. Further, the claims encompass an unlimited amount of additions, therefore, based on the modifications contemplated the claimed protein could have no function or a different function. The specification does not describe properties of the claimed fragment, such as size; or demonstrate any such fragment retaining the activity of the native protein. Note also that claim 2 recites the language "part or all of either of these sequences" and the claim does not recite a functional limitation, therefore, the claimed protein once modified may not be biologically active or may not retain. Additionally, claim 13 recites "a host cell that comprises an expression system which is capable of expressing the protein...". The language "capable of " recited in the claim is not demonstrative of the claimed protein actually being expressed as the term "capable of" means that the event may not occur.

Moreover, claim 13 is directed to a host cell that is capable of expressing a part of the protein and there is no guidance provided as to what part of the protein the host cell is capable of expressing.

The instant specification does not demonstrate or provide guidance as to what the structure of the protein will be once modified or if said protein will be functional or exhibit the same properties or characteristics as the native protein. Additionally, there is no data provided demonstrative of a particular portion of the structure that must be conserved. The art recognizes Filamin A regulates cortical cell migration out of ventricular zone (Nagano et al., Nat. Cell Biol., 2002, July, vol. 4, no.7, pages 495-501). A search of the claimed sequences discloses proteins that are 69.9% or 19% identical to the claimed sequence, however, have a different function and these proteins are encompassed in the claim. For example, HYSEQ INC. disclose a protein that has 73.68% identity to SEQ ID NO:1 and SEQ ID NO:2 (DNA and the encoding the protein, respectively), however, the reference indicates that the protein is useful for treating diseases of the peripheral nervous system, such as neuropathies like Alzheimer's, Parkinson's disease, Huntington's disease, Shy-Drager Syndrome, Amyotrophic lateral sclerosis etc., therefore, the instant claims and specification needs to provide sufficient information regarding the activity to the protein to be altered in the claims. Thus, due to the large quantity of experimentation necessary to generate the infinite number of variants/fragments recited in the claims and possibly screen same for activity and the lack of guidance/direction provided in the instant specification, this is merely an invitation to the skilled artisan to use the current invention as a starting point for further

experimentation. Thus, undue experimentation would be required for a skilled artisan to make and/or use the claimed invention commensurate in scope with the claims.

Predictability of which potential changes can be tolerated in a protein's amino acid sequence and obtain the desired activity requires a knowledge of and guidance with regard to which amino acids in the protein's sequence, if any, are tolerant of modification and which are conserved (for example, expectedly intolerant to modification), and detailed knowledge of the ways in which the protein's structure relates to its function. In addition, one skilled in the art would expect any tolerance to modification for a given protein to diminish with each further and additional modification, for example, multiple substitutions. In this case, the necessary guidance has not been provided in the specification. Therefore, while it is known in the art that many amino acid substitutions are possible in any given protein, the positions within the protein's sequence where such amino acid substitutions can be made with a reasonable expectation of success are limited, as certain positions in the sequence are critical to the protein's structure/function relationship. It is also known in the art that a single nucleotide or amino acid change or mutation can destroy the function of the biomolecule in many cases. For example, various sites or regions directly involved in binding activity and in providing the correct three-dimensional spatial orientation of binding and active sites can be affected (see Wells, Biochemistry, vol. 29, pages 8509-8517, 1990). The instant specification provides no guidance/direction as to which regions of the protein would be tolerant of modifications and which would not, and it provides no working examples of any variant sequence that is encompassed by the claims. It is in no way

predictable that randomly selected mutations, such as deletions, substitutions, additions, etc., in the disclosed sequences would result in a protein having activity comparable to the one disclosed. As plural substitutions for example are introduced, their interactions with each other and their effects on the structure and function of the protein is unpredictable. The skilled artisan would recognize the high degree of unpredictability that all the fragments/variants encompassed in the claims would retain the recited function.

The state of the prior art provides evidence for the high degree of unpredictability as stated above. Seffernick et al. (J. Bacteriology, vol. 183, pages 2405-2410, 2001) disclose two polypeptides having 98% sequence identity and 99% sequence identity, differing at only 9 out of 475 amino acids (page 2407, right column, middle and page 2408, Fig. 3). The polypeptides of Seffernick et al. are identical along relatively long stretches of their respective sequences (page 2408, Fig. 3), however, these polypeptides exhibit distinct functions. The modifications exemplified in the Seffernick et al. reference is small compared to those contemplated and encompassed by the claimed invention (see page 21 of the specification and claim 3, for example). Further, Saus et al. (US PGPUB No. 2003010855A1, 2002) disclose a DNA encoding a protein that is 31.72% identical to SEQ ID NO:1, which encodes SEQ ID NO:2 of the instant application, and it is disclosed that the protein is a GIP family protein, proteins with transcription factor activity. The DNA encoding the protein of the Saus et al. reference is encompassed in the claimed limitation of 1 or several deletions, which demonstrates the unpredictability of the fragment.

The specification lacks adequate guidance/direction to enable a skilled artisan to practice the claimed invention commensurate in scope with the claims. Furthermore, while recombinant and mutagenesis techniques are known in the art, it is not routine in the art to screen large numbers of mutated proteins where the expectation of obtaining similar activity is unpredictable based on the instant disclosure. The amino acid sequence of a protein determines its structural and functional properties, and predictability of what mutations can be tolerated in a protein's sequence and result in certain activity, which is very complex, and well outside the realm of routine experimentation, because accurate predictions of a protein's function from mere sequence data are limited, therefore, the general knowledge and skill in the art is not sufficient, thus the specification needs to provide an enabling disclosure.

The working examples provided do not rectify the missing information in the instant specification pertaining to the claimed fragment/variant. The nature and properties of the claims is difficult to ascertain from the examples provided as one of skill in the art would have to engage in undue experimentation to construct the variants/fragments of the claimed invention and examine the same for function.

The specification does not provide support for the broad scope of the claims, which encompass an unspecified amount of fragments. The claims broadly read on any fragment for the given sequences (SEQ ID NOS: 2, 4 and 6). The issue in this case is the breadth of the claims in light of the predictability of the art as determined by the number of working examples, the skill level artisan and the guidance presented in the instant specification and the prior art of record. This make and test position is

inconsistent with the decisions of *In re Fisher*, 427 F.2d 833, 839, 166 USPQ 18, 24 (CCPA 1970) where it is stated that "... scope of claims must bear a reasonable correlation to scope of enablement provided by the specification to persons of ordinary skill in the art...". Without sufficient guidance, determination of having the desired biological characteristics is unpredictable and the experimentation left to those skilled in the art is unnecessarily and improperly extensive and undue. See *In re Wands*, 858 F.2d at 737, 8 USPQ2d at 1404 (Fed. Cir. 1988). Therefore, absent direction/guidance regarding whether the structure of the encoded protein can tolerate the modifications contemplated a non-functional protein may result and one of skill in the art would not be able to practice the claimed invention commensurate in scope with the claims.

Thus, for all these reasons, the specification is not considered to be enabling for one skilled in the art to make and use the claimed invention as the amount of experimentation required is undue, due to the broad scope of the claims, the lack of guidance and working examples provided in the specification and the high degree of unpredictability as evidenced by the state of the prior art, attempting to construct and test variants of the claimed invention would constitute undue experimentation. Making and testing the infinite number of possible fragments to find one that functions as described is undue experimentation. Therefore, applicants have not provided sufficient guidance to enable one of skill in the art to make and use the claimed invention in a manner that reasonably correlates with the scope of the claims, to be considered enabling.

The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter, which the applicant regards as his invention.

9. Claims 1-3 are rejected under 35 U.S.C. 112, second paragraph, as failing to set forth the subject matter, which applicant (s) regard as their invention.

Claim 1 is indefinite for the recitation of one or several amino acids are deleted because there is no upper limit, therefore, the entire sequence could be deleted. The dependent claims hereto are also included in this rejection as they do not rectify the deficiency.

Claim 3 is indefinite for the recitation of "e.g." in association with the hybridization conditions because this is not limiting. It is well known in the art that hybridization conditions can vary thus, if an example is provided and not the actual conditions applicant intends, the metes and bounds of the claim is unclear. The claim is also confusing, note that line three of the claim has a period following the pH and another sentence that cannot stand alone. It is suggested that the claim is amended to recite the hybridization conditions that is intended for the claimed invention and delete the period. Note also the phrase "stringent conditions" is recited twice in line two of the claim.

Claim Rejections - 35 USC § 102

The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless –

(b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.

10. Claims 1-3 and 13 are rejected under 35 U.S.C. 102(b) as being anticipated by Yen et al. (U.S. Patent No. 5599919, February 4, 1997), based on the broad recitation of one or several amino acids are deleted, substituted or added; a sequence comprising part or all of either of the sequences.

Yen et al. disclose a DNA encoding a protein which is 20.6% identical to the claimed sequences set forth in SEQ ID NOS: 1 and 2, therefore, which anticipates claims 1-2. In addition, the patent teaches expression in a host cell (claim 13, see column 7). As the referenced sequence has 20.6% sequence identity to the claimed sequences, the referenced DNA would hybridize to the claimed sequence. The functional property recited in the claims although not taught by the reference would be construed as an inherent property. Therefore, the limitations of the claims are met by this reference.

Conclusion

11. No claims are presently allowable.

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Any inquiry concerning this communication or earlier communications from the examiner should be directed to Hope A. Robinson whose telephone number is 571-272-0957. The examiner can normally be reached on Monday-Friday.

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Jon P. Weber, can be reached at (571) 272-0925. The fax phone number for the organization where this application or proceeding is assigned is 571-273-8300.

Information regarding the status of an application may be obtained from the Patent Application Information Retrieval (PAIR) system. Status information for published applications may be obtained from either Private PAIR or Public PAIR. Status information for unpublished applications is available through Private PAIR only. For more information about the PAIR system, see <http://pair-direct.uspto.gov>. Should you have questions on access to the Private PAIR system, contact the Electronic Business Center (EBC) at 866-217-9197 (toll-free).

Hope Robinson, MS

Patent Examiner

4/14/05

| | | | |
|----|---------------------------|------------------------------------------------------------------------|------|
| Qy | 1875 | GGCATAGAGGGTAGAAAGGAATAACCGAGGTAGGTCTGAGTC | 1934 |
| Db | 2955 | LGCAAGGCGCCAGGCGGCTTCTGAGGAGGCGCTTGCGAC | 3074 |
| Qy | 961 | SerGlnLysProLyserSerIleAspProThrLeuGlyProIleAlaMetSerProVal | 980 |
| Db | 601 | GlyIleGluGluIleGluIleArgGluIleAsnArgGlyIleArgSerCysLysGlySerGluPhe | 620 |
| Qy | 3015 | ACGATTACTATTCAGAGAGAGGCCAGGGACTAACGCTGAAATCGAGAGACTGAAAGAAA | 1994 |
| Db | 1935 | ACCTGCCGAGACAATAAGATCAGAGAAGCTAACGCTGAAATCGAGAGACTGAAAGAAA | 1994 |
| Qy | 621 | ThrCysProGluAspAsnLysIleArgGluIleThrLeuGluIleArgGlyIleArgSerIleAsn | 640 |
| Db | 641 | ArgLeuGlnGlnIleLeuGluValValGluGlyAspLeuThrLeuGlyAspGluLeuLys | 640 |
| Qy | 1995 | CGGCTCCAGCAGTTGGAGGTGGAGGGACTGTGAGAGAGGAGGAATAGCAACTTCCCTCCAGCAGCTC | 2054 |
| Db | 661 | GlnLeuGluGlnLysPheArgThrGluGlnAspLysAsnProLeuSerGlnGlnLeu | 680 |
| Qy | 2055 | CAGTGGAGCAGAAGTTCAGAACCCAGGAGGAGATAAGCCAACCTCCCTCCAGCAGCTC | 2114 |
| Db | 681 | GluGlyIleLysHisAsnMetAlaLysLysAlaIleGluIleGlyIleAsnProLeuSerGlnGlnLeu | 700 |
| Qy | 2115 | GAGGAATCACACCAAATGGCCAGCACAGCCATAGAGAAGGGAGGGCTAAGTGTGATTACAG | 2174 |
| Db | 681 | GluGlyIleLysHisAsnMetAlaLysLysAlaIleGluIleGlyIleAsnProLeuSerGlnGlnLeu | 700 |
| Qy | 2175 | CAGGAAGCCGAACTGCCCACAGGTTGGCTGGAGGAGCTAAGTGTGATTACAG | 2234 |
| Db | 701 | GlnGluIalGluLeuArgHisArgPheArgThrGluIleAlaLysSerArgAspLeuGln | 720 |
| Qy | 2235 | TCTCAGCTCAAGTCAGGAGAGATCCAGGCTGATGAAACAAGGAAGCAGCTG | 2294 |
| Db | 721 | AlaGluValGlnIleAlaLysLysGluLysIleHisGluLeuMetAlaLysGluAspGlnLeu | 740 |
| Qy | 2295 | TCTCAGCTCAAGTCAGGAGAGATCCAGGCTGATGAAACAAGGAAGAAGCTG | 2354 |
| Db | 741 | SerGlnLeuGlnValAspTyrSerValLeuGlnGlnArgPheMetGluGluGluThrLys | 760 |
| Qy | 2355 | ACAAAGAACATGGGAGGGAGCTCTAACATGACCAGGCTAGACCTTCAACCC | 2414 |
| Db | 761 | AsnLysAsnMetGlyArgGluValLeuAsnLeuThrLysGluIleGluLeuSerAlaLysBrg | 780 |
| Qy | 2415 | TACAGCCGAGCTCTAGGAGGAGGGAGGAGGAGGTTGGCCCTGGCC | 2474 |
| Db | 781 | TyrSerArgAlaLeuArgProSerGlyAsnGlyArgMetValAspValProValAla | 800 |
| Qy | 2475 | TCCACTGGGGAGCCGAGGGCTGTCGGGGATCTGGGGAGGACCCGCT | 2534 |
| Db | 801 | SerThrGlyValGlnThrGluIleValCysGlyAspAlaIleGluGluIleThrProAla | 820 |
| Qy | 2535 | GTGTCATGCCAACATCCCTCAGGAGAAATCACATCATGAGTAATCTGGACAGGT | 2594 |
| Db | 821 | ValPheIleArgLysSerPheGlnGluAsnHisIleMetSerAsnLeuArgGlnVal | 840 |
| Qy | 2595 | GGCCTGAGAACCCATGGAACGGTCTCGTCCACAGGTATCCCCCAGGCAAT | 2654 |
| Db | 841 | GlyLeuLysLysProMetGluArgSerSerValLeuAspArgTyrProProAlaAlaAsn | 860 |
| Qy | 2655 | GAGCTCACCAGTGGAGCTGCTGGATTCCTGGATGAGAAAGAGAACGGTCTTCC | 2714 |
| Db | 861 | GluleuThrMetArgLysSerTrpIleProTrpMetArgLysArgGluAsnGlyProSer | 880 |
| Qy | 2715 | ACTCCGCAGGAGAACGGCCAAACAGGGTCAGGGCACCCGGGAGCTGCTC | 2774 |
| Db | 881 | ThrProGlnGluLysGlyProArgProAsnGlnGlyIleProGlyGluLeuVal | 900 |
| Qy | 2775 | CTAGCACCAGGAGGAGCTGGATTCCTGGATGAGAAAGAGAACGGTCTTCC | 2834 |
| Db | 901 | LeuIleProLysGlyIleArgGlyProIleLeuIleAspLeuIleAspAspHisGluSer | 920 |
| Qy | 2835 | ACTGCCACCCCTGGAGATCACAGCCCACATCTGAAGAGTTCTAGTACACCGTC | 2894 |
| Db | 921 | ThraIleThrLeuGluIleThrSerProThrSerGluGluIlePheSerSerThrThrVal | 940 |
| Qy | 2895 | ATTCCTACCTAGGCAACCAAGAAAGATAACCATATTCCATCACCAATGTCATG | 2954 |
| Db | 941 | IleProThrLeuGlyAsnGlnLysProArgIleThrIleProSerProAsnValMet | 960 |
| Qy | 941 | Homo sapiens. | |
| Qy | W0200153312-A1. | | |
| DB | RESULT 2 | | |
| ID | AAM40016 | standard; protein; 1213 AA. | |
| XX | | | |
| AC | AAM40016; | | |
| DT | 22-OCT-2001 (First entry) | | |
| XX | | | |
| DR | | Human polypeptide SEQ ID NO 3161. | |
| XX | | | |
| KW | | Human; nootropic; immunosuppressant; cytostatic; gene therapy; cancer; | |
| KW | | peripheral nervous system; neuropathy; central nervous system; CNS; | |
| KW | | Alzheimer's; Parkinson's disease; Huntington's disease; haemostatic; | |
| KW | | amyotrophic lateral sclerosis; Shy-Drager Syndrome; chemotactic; | |
| KW | | chemokinetic; thromolytic; drug screening; arthritis; inflammation; | |
| KW | | leukaemia. | |

PP 26-JUL-2001.
 PR XX
 PR XX 23-DEC-1999; 99US-00471275.
 PR 25-APR-2000; 2000US-00488725.
 PR 20-JUN-2000; 2000US-00552317.
 PR 19-JUL-2000; 2000US-00598042.
 PR 03-AUG-2000; 2000US-00620312.
 PR 14-SBP-2000; 2000US-00653450.
 PR 19-OCT-2000; 2000US-00662191.
 PR 29-NOV-2000; 2000US-00693036.
 PA XX
 PA (HYSEB-) HYSEQ INC.
 PI XX
 PI Tang YT, Liu C, Abundi V, Chen R, Ma Y, Qian XB, Ren F, Wang D, Wang J, Wang Z, Wehrman T, Xu C, Xue AJ, Yang Y, Zhang J, Zhao QA;
 PI Zhou P, Goodrich R, Drmanac RT;
 DR WPI; 2001-442253/47.
 PT N-PSDB-PAT5972.
 PT Novel nucleic acids and polypeptides, useful for treating disorders such as central nervous system injuries.
 PS Example 4; SEQ ID NO 3161; 10078pp; English.
 CC The invention relates to human nucleic acids (AT57798-AA161369) and the encoded polypeptides (AA38642-AA42213) with nootropic, immunosuppressant and cytostatic activity. The polymucleotides are useful in gene therapy. A composition containing a polypeptide or polynucleotide of the invention may be used to treat diseases of the peripheral nervous system, such as peripheral neuropathy and localised neuropathies and central nervous system diseases, such as Alzheimer's, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and Shy-Drager syndrome. Other uses include the utilisation of the activities such as: Immune system suppression, Activin/inhibin activity, chemotactic/chemokinetic activity, haemostatic and thrombolytic activity, cancer diagnosis and therapy, drug screening, assays for receptor activity, arthritis and inflammation, leukaemias and C.N.S disorders. Note: The sequence data for this patent did not form part of the printed specification
 CQ Sequence 1213 AA;
 CQ Alignment Scores:
 CQ red. No.: 0
 CQ core: 5696.50
 CQ Percent Similarity: 96.54%
 CQ Local Similarity: 93.49%
 CQ Query Match: 73.68%
 CQ 4
 CQ Gaps: 1
 CQ Sequence 1213 AA;
 CQ 75 ATGAGATCAGGAATCAAGGTGGAGAAGTTCACTAACGGCATGTCCTGCCAAG
 CQ 1 MetargserArgasnGinglygluseralaseraspGlyHisIleSerCysProly 20
 CQ 135 TCCTCCATCATCAGCAGTGTGTTAGGGCCCTCAGAAGATGCA--AAAAGAAC
 CQ 21 ProserileleGlyAsnAlaGlyGlulyLysSerLeuserGlyHisIleSerCysProly 20
 CQ 192 AGGCCAATCGGAGGAGGAGGATCTAGGCCTCCGAACATCAAAGGCCCTCAA
 CQ 41 LysserasnarglysgluaspAspValMetAlaSerGlyThrValLysLysLys 40
 CQ 252 CCATCTGGAGAGACTAAGACTAGACTGAGTGTGAGATGAGGAGGAGCTCCAG
 CQ 61 ThrserGlyGluCysLysValLysThrLysSerLeuLysLysLysLys 60
 CQ 312 ATCCAGCTCTTAGTATCATGGAAGGAGGAGGAGCTGAGGAGGAGGAGCT 311
 CQ 421 LeuargleugluValGlyLysLysLysLysLysLysLysLysLys 80
 DB 81 IieGinieLeuseRileMetGluGlyGluGlnAlaArgGluAspValLeuHisMet 100
 DB 372 CTGAGGACAGAGAAAACCAAGCCGAGTTCTGGAGCACACTATGGATCTGCGAGACT 431
 DB 101 LeuLysThrGlyLysThrLysProGluValLeuGluAlaHisTyrglySerAlaGluPro 120
 DB 432 GAGAAAGTGCTCGGCTCTCACCGAGATGCCATCCCTGCTCAAGAGAAGTCATAGGA 491
 DB 121 GlyLysValLeuArgValLeuHisArgAspAlaIleLeuAlaGlyLysSerIleGly 140
 DB 492 GAAGACGCTTATGAGAACCTATCAGAGCTGAGCACAGACTGGAGAAAGCAGAGGAG 551
 DB 141 GluAspValTyrGlyLysProIleSerGluLeuAspArgLeuGluGlyGlnlysGlu 160
 DB 552 ACGTACGGCCATCTAGAGCAGCTGCTGCTGGCTGAGAAGTGTCAACAGGGCACCGTG 611
 DB 161 ThryTyrglyArgMetLeuGluGlyLysGluLysGlyLysGlyLysGlyLysGly 180
 DB 612 TAGGAGCTGGAGAACGAGACAAGCACACTGACTACATGAAACAAGGGAGGACTTC 671
 DB 181 TyrGlyLeuGluAsnGlyLysLysLysLysLysLysLysLysLysLysLysLys 200
 DB 672 ACCAACCTCTGGAGCAGGAGGAGGAGGCTGAAAGCTCTGAAACAAAGAAAGCT 731
 DB 201 ThrAsnLeuIeuGluGlyLysGlyLysLysLysLysLysLysLysLysLysLysLys 220
 DB 732 TACCAAGCCGAAAGAAACGAAACGCTTAAGGGCTAACAAACTTCGAGATGAGCT 791
 DB 221 TyrglyAlaArgLysGlyLysGluAsnAlaLysGlyLysGlyLysGlyLysGly 240
 DB 792 CTGAGGCTCAACTCCTTCGCCCCTCAGTGTGGACGGAGGAGGAGCTGAGCAA 851
 DB 241 ValIysLysLeuIeuGluGlyLysGlyLysGlyLysGlyLysGlyLysGlyLysGly 260
 DB 852 CTGAGGCTCAGACTCAGAACTCAGAACGAGGAGGAGGAGGAGGAGGAGGAGA 911
 DB 261 LeuGlyLeuLysSerPheAlaLeuMetLeuValAspGluArgGlyLysLysLys 280
 DB 912 AACCTCAAAGCGGTCACTACAAATCCAAGGAAGACCAGGCCAGAACGCTGCTCAAGTAGAA 971
 DB 281 LysLeuLysAlaIleThrSerLysSerLysLysLysLysLysLysLysLysLys 300
 DB 972 GTGCACTTCGAAACACAGGCCTCAGGTTTCCAGGAGGAGATGAAACGCCAA 1031
 DB 301 ValAspPheGlyLysLysAlaSerArgPheSerGlyLysLysLysLysLysLysLys 320
 DB 1032 TTGGCGAATCAAAGATCTCACACCGCAACTTCGACTCAACTGGTTGCTTATCGCAA 1091
 DB 321 LeuAlaAsnGlyGluSerHisIleSerGlyLysLysLysLysLysLysLysLysLys 340
 DB 1092 AGGATTGAGGAGCTGGAGAGACCAATAAGCCTTCAGAGGAGGAGGAGCTCCAG 1151
 DB 341 ArgIleGlyIleLeuGlyLysLysAsnLeuLysSerIleSerLysLysLysLysLys 360
 DB 1152 GAGCTGAGAGAGAAATGCCAAACGGGAACTTCAGTCTCATGGGGAGTG 1211
 DB 361 GluLeuArgAspLysIleAlaLysSerIleGlyLysGlyIleSerSerIleLeuMetAlaGlu 380
 DB 1212 GAGCTGAGAGAGAAATGCCAAACGGGAACTTCAGTCTCATGGGGAGTG 1271
 DB 381 GluAsnLeuArgLysGlyLysValLeuGlyLysLysLysLysLysLysLysLysLys 400
 DB 1272 GAGCTGAGAGAGAAATGCCAAACGGGAACTTCAGTCTCATGGGGAGTG 1331
 DB 401 GluAsnGlyLysSerIleGlyLysGlyLysLysLysLysLysLysLysLysLysLysLys 420
 DB 1332 CTTAGACTAGTGGAGAGCTGAGAGAGGAGCTGAGCTGGAGAGGAGGAGA 1391
 DB 421 LeuArgLeuGlyLysValGlyLysLysLysLysLysLysLysLysLysLysLysLysLys 440

QY 3612 CTGACCAATTCCAGCCTCGAGCTGAGACTCAGTCTATGAAATAGACCTGAAGAATCT 3671
Db 1181 LeuthrLysPheGluProArgAlaGluThrGinSerMetLysIleLeuIlySlySer 1200

QY 3672 GCAGCCAGCAGCACTGCCTCTGGAGGGAGGGC 3710
Db 1201 AlaLaserSerThrSerLeuGlyGlyGlyLysGly 1213

RESULT 3
ID ABP97031
ID ABP97031 standard; protein; 1213 AA.

DB Human L-PILIP protein SEQ ID NO:6.

XX L-PILIP; S-PILIP; filamin-interacting protein; cell migration; cell death; cytostatic; neuroprotective; immunosuppressive; cancer; KW tumour metastasis; transplantation therapy.

XX Homo sapiens.

XX WO2003018804-A1.

XX 06-MAR-2003.

XX PR 29-JUL-2002; 2002WO-JP007676.

XX PA (NISC-) JAPAN SCI & TECHNOLOGY CORP.

XX PI sato M., Nagano T;

XX DR WPI; 2003-268423/26.

XX N-PSDB; ACC45356.

PT Proteins controlling cell migration and cell death and their encoded DNAs, applicable in developing drugs for treating or suppressing cancer or tumor metastasis or as regulators of cell migration for transplantation.

XX PS Claim 7; Page 82-88; 96pp; Japanese.

CC The present sequence represents human L-PILIP which is a filamin-interacting protein. L-PILIP has a function of controlling cell migration and cell death. L-PILIP has cytostatic, neuroprotective and immunosuppressive activities. The L-PILIP protein can be used for controlling cell migration and cell death, which is applicable in developing drugs for treating or suppressing cancer or tumour metastasis or as regulators of cell migration for transplantation therapy, and also for controlling the mobility and cell death of nerve cells, promoting decomposition of the actin-binding protein e.g. filamin-interacting protein in the treatment of pre-ventricular nodular heterotopia

XX Sequence 1213 AA;

Alignment Scores:
Pred. No.: 0 Length: 1213
Score: 5696.50 Matches: 1134
Percent Similarity: 96.54% Conservative: 37
Best Local Similarity: 93.49% Mismatches: 41
Query Match: 73.68% Indels: 1
DB: 6 Gaps: 1

US-10-788-793-1 (1-4364) x ABP97031 (1-1213)

QY 75 ATGAGATCAGGAATCAAGTGGAGAACTCATCTAACGGGCAATGTCCTGCCCAAG 134
Db 1 MetArgSerArgArgGlyGlyGluSerAlaSerAspGlyHisIleSerCysProLys 20

QY 135 TCCCTCCATCATCACAGCTGATCGTGGTAAGGCCCCCTCAGAGATGCA---AAAAGAAC 191
Db 21 ProSerLeuIleGlyAsnAlaGlyGluIlySlySerIleSerGluAspAlaLysIleIlySlySer 40

QY 192 AAGGCCATCGGAGGAGGATGTCTGGCTCCGCACTATCAAAGGCACCTCAA 251
Db 41 LysSerAsnArgLysGluAspAspValMetAlaSerGlyThrValLysSerGlyIleIlySlySer 60

QY 252 CCATCTGGAGAGAGGAAAGACTAAGAAGTCTGAGAGACTGCTGGAGTTATCCAGGAGGACCTC 311
Db 81 IleGlnLeuLeuSerIleMetGluIlyGluLeuGlnAlaArgGluAspValLeuIleIlySlySer 80

QY 312 ATCCAGCTCTGAGTATCATGGAGGGAGTTCAGGCTCGAGAGATGTCATCCACATG 371
Db 101 LeuIlySlySerIleMetGluIlyGluLeuGlnAlaArgGluAspValLeuIleIlySlySer 100

QY 432 GAGAAAGTGCTTCGGGCTCTGCAACGAGATGCCATCTGCTCAAGAGAAGTCCATAGGA 491
Db 121 GluIlySvalLeuArgValLeuIleAspAlaLeuIleIlyGluIlySerIleIlyGlu 140

QY 492 GAAGACGCTATGAGAACCTATCTCAGAGCTGGACAGACTGGAGGAAAGGAGGAG 551
Db 141 GluAspValTyRArgMetIleGluIlySlyProIleSerGluLeuAspPargLeuGluIlySlyGlu 160

QY 552 ACGTACCGCCATGCTAGGCCAGCTGCTGCTGAGTGGAGAGTCAGGGCACCGTG 611
Db 161 ThrTyRArgArgMetIleGluIlySlyProIleSerGluLeuAspPargLeuGluIlySlyGlu 180

QY 612 TAGAGCTGGAAACGAGGACAAGCACACTGACTACATGAAACAGGCAACGCTC 671
Db 181 TyRgluIleGluIaAsnGluIlySlyIleIlySlyIleAspPtyrMetAsnLysSerAspPhe 200

QY 672 ACCAACCTGCTGGAGGGAGGAGAGGTCAAAGCTCTGACAAAGGAGCTC 731
Db 201 ThrAsnIleLeuIleGluGlnIleGluIlySlyIleAspPtyrMetAsnLysSerAspPhe 220

QY 732 TACCAAGCCCCAAAGAAAGGAAACGCTAACAAACTCGAGATGCACTCGAGCT 791
Db 221 TyRgluIaArgLysGluIlySlyIleAspPhe 240

QY 792 GTGAACTCAAGTCTCCCTCATGTTGCTGAGGAGATGCACTCGAGCT 851
Db 241 ValIlySlyIleIlySlySerPheAlaLeuMetIleIvaIAspGluArgGlnMetThrIleGluIle 260

QY 852 CTGGACCTGAGACTCAGAAAGTCCAGGACCTACTCAGAACCTGAGGAGGAGAA 911
Db 261 LeuGlyIleGluIlySlySerGlnIlySlyValGlnAspIleThrGlnIlyIleIlySlyLeuIleGluIle 280

QY 912 AACTCAAGGGTCACTTACAATCCAGGAGACGCCAGAGCTCTCAAGTAGAA 971
Db 281 LysLeuIlySlyIleIlySlySerIleIlySlyGluAspArgGlnIlyIleIlySlyLeuIleIlySlyLeuIle 300

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QY 1032 TTGGCGAATCAGAACTCTCACAAACGGCAACTCTGACTCACACTGGCTTATCGAA 1091
Db 321 LeuIalaAsnGlyIleGluIlySlySerIleIlySlySerGlnIlyIleIlySly 340

QY 1092 AGGATTGAGGAGCTGGAGAGACCAATAAGGCTTCAGAGGGAGAGCTCCAG 1151
Db 341 ArgIleGluIleGluIlySlyIleIlySlySerIleIlySlySerIleIlySly 360

QY 1152 GACCTGAGGAGAAATGCCAAGGGAAATGTTGAAACTCCAGTCTCATGGCGAAGTG 1211
Db 361 GluIleIlySlyIleIlySlyIleIlySlyIleIlySlyIleIlySlySerIleIlySly 380

QY 1212 GAGCTCTGCAAGGCCGCTGCTGAGTGGAGGCAAGGATGAAGAGATCACGAAGACC 1271

2409 AAGGGCTACAGCCGAGCTCTAGGCCAGTGGAACGGCGAAGGATgtggaaacgtgcct 246
 681 LysargtryrSerArgAlaLeuargProSerValAsnGlyArgArgMetValAspValPro 700
 701 ValThrSerThrGlyValGlnThrAspAlaValSerGlyGluAlaAlaGluGluThr 720
 721 ProAlaValPheIleArgLysSerProGlyGluGluAsnIleMetSerAsnLeuarg 740
 740 GlnGlyGluLysProValGluArgSerSerValLeuAspArgTyxProProAla 760
 761 AlaAsnGluLeuThrMetArgLysSerTrpIleProTrpMetArgLysArgGluAsnGly 780
 780 CCTTCCACTCCGAGGAGAAAGGCCAGGCCAAACCAAGGGCACCCGGGAG 2768
 781 ProSerIleThrGlnGlyProArgThrAsnSerSerProGlyHisProGlyGlu 800
 782 CTTGGATTCAGGAGAAAGGCCAGGCCAAACCAAGGGCACCCGGGAG 2768
 783 ValValLeuSerProLysGlnGlyGinProLeuHisIleArgValThrProAspHisGlu 820
 784 AACAGCACTGCCACCCGAGATCACAGCCCACATCTGAAGAGTTTCTTAGTAC 2888
 821 AsnSerThrAlaThrLeuGluIleThrSerProThrSerGluGluPhePheSerSerThr 840
 840 ACCGTCAATTACCTAGGCAACCAGAAACCAATAACCATATCCATCACCAAT 2948
 841 ThrValIleProThrLeuGlyAsnGlnIleProArgIleThrIleProSerProAsn 860
 860 GTCATGTCGAAAGCCAAAGTGCAGATCCTACTCTGGCCAGAACGGCATGTCC 3008
 861 ValMetProGlnIlySglNlysSerGlyAspThrThrLeuGlyProGluArgAlaMetSer 880
 880 CCTGTCACGATTACTATTCAGAGAGAACCGGAAAGCTGAAAGGAGCCCTT 3068
 881 ProValThrIleThrThrPheSerArgGluLysThrProGluSerGlyArgGlyAlaPhe 900
 900 GCCGACAGGCCATCCCCATCCAATCATGACGGCTCACATCTGCAGCTCCACT 3128
 901 AlaAspArgProThrSerProIleGlnIleMetThrValSerThrSerAlaAlaProAla 920
 920 GAATCGCTGCTCTGAATCTCAGGAAGTGCCTATGGGAAGGACTATCCTCAAGTC 3188
 921 GluIleAlaValSerProGluUserProGlnGluMetProMetGlyArgThrIleLeuLysVal 940
 940 ACCCCGAAACAACTGTTCCAGCCCCGTGCGAAAGTACAACCTCAAATGCTTAATC 3248
 941 ThrProGluIlySglNthrvAlProThrProValArglyStyAsnSerAsnAlaAsnIle 960
 960 ATCACCACCGAAAGCAATAATCACATTCACTGAGGTTCTCAAGTAAAGCATCT 3308
 961 IleThrThrGluAspAsnLysIleHisIleHisIleLeuGlySerGlnPhelySargSerPro 980
 980 GGGCCTGCCGCTGAAGGCCAGTTACCGTccGcGtcaACGTGACAGCG 3368.
 981 GlyThrSerGlyGluLysValSerProValIleThrValArgProValAsnValIleAla 1000
 1000 CCCGGTGTAGCAAGTGACCGCACTATAACTAACCCTGACAAGTCATCCACA 3428
 1021 ProGlyAlaSerLysValThrSerThrIleThrProValThrSerSerAla 1040

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Alignment scores:
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 Score: 2452.50
Percent similarity: 64.40%
Best Local Similarity: 46.17%
Query Match: 31.72%
Length: 113
Matches: 542
Conservative: 214
Mismatches: 337
Indels: 81
Gaps: 5

US-10-788-793-1 (1=436A) X MS-10-300-8E1 14 11 1963

75 ATGAGATCACGAATCAAGTCGAGAAAGTTCATCTAACGGCATGTCCTGCCCAAG 13
 1 MetargSerarg---GlySeraspThrGluGlySerAlaGlnLysAspPheProArg 18
 135 TCCTCCATCATCAGCACTGATGGTGTAAAGGCCCTCAGAAGATGCCAAAGAACAG 19
 19 Histhr-----LysGlyHisSerPheGlnGlyProLysAsnMet 31
 195 GCCAAATCGGAAGGGAGGAG--GATGTCATGGCTCCGAACTATCAAAGCACCTCAA 251
 32 LysHisArgGlnGlnAspLysAspSerProSerProAspVal-----IleLeu 48
 252 CCATCTGGAGAAAGTGAGAA-----AAGACTAAGAAGTCTGGAGTTATCC 299
 49 ProCysProLysAlaGluLysProHisSerGlyAsnGlyHisGlnAlaGluAspLeuSer 68
 300 AAGGAGGACCTCATCCAGCTCCTGAGTATCATGGAGGGAGTTGCAGGCTCGAGAAAGAT 359
 69 ArgAspAspPheLeuPheLeuSerTleLeuGluGlyGluLeuGlnAlaArgAspGlu 88
 360 GTCATCCACATGCTGAGGACAGAGAAACCAAGCCCCGAGGTTCTGGAGGACACATATGGA 419
 89 ValIleGlyIleLeuLysAlaGluLysMetAspLeuAlaLeuGluAlaGlnArgLysSer 108
 420 TCTGGAGAACCTGAGAAAGTGCTTCGGGTCTGCCGAGATGCCCATCCTGCTCAAGAG 479
 109 PheValIleThrProLysLysValIleGluAlaLeuGlnArgAspAlaPheGlnAlaLysSer 128

CURRENT APPLICATION NUMBER: US/10/309,851
 CURRENT FILING DATE: 2002-12-04
 NUMBER OF SEQ ID NOS: 38
 SOFTWARE: PatentIn version 3.1
 SEQ ID NO 16
 LENGTH: 1133
 TYPE: PRT
 ORGANISM: Homo sapiens
 US-10-309-851-16

Alignment Scores:
 Pred. No.: 5.87e-139 Length: 1133
 Score: 2444.50 Matches: 541
 Percent Similarity: 64.31% Conservative: 214
 Best Local Similarity: 46.08% Mismatches: 338
 Query Match: 31.62% Index: 81
 DB: 14 Gaps: 19

US-10-788-793-1 (1-4364) x US-10-309-851-16 (1-1133)

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QY .135 TCTTCCATCATCAGCGAGTGATGGGCTTAAGGGCCCTCAGAAGATGCAAAGAAC 194
Db 19 HisThr-----LysGlyHisSerPheGlnGlyProAsnMet 31
QY 195 GCAAATGGAGGGAG---GATGTCATGGCTCCGGAACTATCMAAGCACCTCAA 251
Db 32 LysHisArgGlnAspLysAspSerProSerGluSerAspVal-----Ileu 48
QY 252 CCATCTGGAGAAAGTGAGAAA-----AAGACTAAGAAGTCGTGGAGTATCC 299
Db 49 ProCysProLysSerGlySerLeuThrProGluArgThrMetSerProIleGlnVal 1004
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Db 69 ArgAspAspLysSerGlySerAlaIleuSerIleuGluGlyGluGlnAlaArgAspGlu 88
QY 3159 GTGCCCTATGGAGGACTATCCTCAAAGTCACCCGGAAACAACACTGTCCAGCCCC 3218
Db 105 IleuAlaValThrGlySerIleuSerProGluGlySerIleu 1024
QY 3219 GRGCGGAAGTACAATCTAACTCTAAATTCACATTCTGAGGAGAAAC 3278
Db 1045 GluArgSerAsnSerAsnSerSerValIleuThrGluAspAlaLysIleu 1064
QY 3279 CACCTGGAGTCTCAGTTAACGGATCTCTGGGCTGAGGCTTACAGGCAGTCCT 3338
Db 1065 HisBleuGlySerProTyroMetGlnAla-----ValIleuSerProValArgProAla 1081
QY 3339 ATCACCGCTCCGGCTGTCAACGGAAAGGGGTTCTACAGGCACAGTCCT 3398
Db 1082 SerProSerAlaProLeuGlnAspSerArgThrGlyIleuLeuAsn 1101
QY 3399 CGCTCTCCCAGGAACCACTCTCTAACAGACCCGGTCTAGCAAGTGGACCACTA 3458
Db 1102 LysThr-----ThrAsnLysValThrSerSerIleu 1111
QY 3459 ACTATAACCCGGTACACGGTCACTCACAGGAGAACCAA 3500
Db 1112 ThrIleuThrProThrAlaThrProLeuProArgGlnSerGln 1125

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RESULT 3

US-10-309-851-16

; Sequence 16, Application US/10309851

; Publication No. US20030108554A1

; GENERAL INFORMATION:

; APPLICANT: Revert-Ros, Francisco

; TITLE OF INVENTION: GIPs, a Family of Polypeptides with Transcription Factor Activity

; FILE REFERENCE: 98,723-F-US

QY 720 CAGAGAAAGCTTACCAAGCCGAAAGAAAGGAACGCTAACAACTT 779
 Db 209 GlnGluIleLysSerGlnGluGluIleGluGlySerValThrThrIleu 228
 QY 780 CGAGATGAGCTTGAGCTCAAGTCCTCGGCTCATGTTGGAGCAGGAG 839
 Db 229 LysGluGluLeuThrLysSerPheAlaLeuMetValValAspGluGlnArg 248
 QY 840 CACATCGAGCACTGGGCTGAGTCAGAAAGTCAGGACCTCACTCGAGCTGAG 899

Db 1964 GluLeuGluLeuThrLysMetAspLysMetSerPheValGlyLysValAsnLysMetThr 1983
 Qy 2172 AGCCAGAACCGAACTGGACACAGGRTCTGGCTGGAGGGCTAAACTCGTGATT 2231
 Db 1984 AlalysGlutThrGluLeuGlnArgGluMetHisGluMetAlaGlnLysThrAlaGluLeu 2003
 Qy 2232 CAGGCCAGGTCAAGCTCAAGGAGAAG 2267
 Db 2004 GlnGluGluLeuSerGlyLysGluLysAsnArgLeuAlaGlyGluLeuGlnLeuLeuGlu 2023
 Qy 2268 GAGCTGATGAACAAGGAGACCAGCTGTCAGTCAAAGTCGACTATTCAGTCAG 2327
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 Qy 2328 CAAGAGTT-----ATGGAAGAAGAACTAAGAACAGAACATGGGAGGGAG 2375
 Db 2044 LysSerLeuAspCysMethIleLysAspGlnValGluLysGluLysGluLeuLys 2083
 Qy 2376 GTCCCTCAATCTGACCAAGGACCTAGAGCTTAGGCTTCAGGCTCACAGCCAGCTCTCAGGCCG 2435
 Db 2064 IleAlaGluThrGlyGlnLeuArgLeuHisGluAlaIleLysLysHisGlnAlaLeuLeu 2103
 Qy 2436 AGTGGGAACCGGGCGAAGGGTAGGGTGGGACGAGCTCCACTGGGGTCAGACCG 2495
 Db 2084 AspPheAsnLysGlnTyRgluValGluIleGlnLysGluLeuLys 2120
 Qy 2496 GCGGRTGCGGGATGCTGGAGGAGACGGAGACCCGAGCTGTTCAATCCTTC 2555
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 Qy 2556 CAGGAGGAA-----AATCACATC----- 2573
 Db 2121 LysGluGluLeuAsnSerSerLysAlaThrThrGlnIleLeuGluGluLeuLys 2140
 Qy 2574 -----ATGAGTAATCTTGACAGGTGAC-----CTGAAGAACCATGGAACGGTCC--- 2621
 Db 2141 ThrLysMetAspAsnLeuLysTyrValAsnGlnLeuLysGluAsnGlu 2160
 Qy 2622 TCGGTCTCGACAGGTATCCCCCAGCGCGAATGAGCTCACCATGAGGAAGTCTGGATT 2681
 Db 2161 GlyLysMetIleLeuLeuIleLysSerCysLysGlnLeuGluGluLysGlu----- 2178
 Qy 2682 CCTTGATGAGAAAGAGAACGGTCCACTCCGAGGAGAACGGCCAGGCCA 2741
 Db 2179 ---IleLeuGlnLysGluLeuSerGlnIleGlnIleAlaIleGlnLys----- 2193
 Qy 2742 AACCAAGGTGCAGGCACCCGGGAGCTGGTCTTACCAAGGAGCCAGCCCCTA 2801
 Db 2194 ----- 2201 AspThrLysValAspGluLeuThrThrGluLysGluLeuLysGluThrLeuGlu 2220
 Qy 2802 CACATCCGRTG-----ACACCAAGATCATGAGAACGAGCTGCCACCCGGAGTC 2852
 Db 2201 AspThrLysValAspGluLeuThrThrGluLysGluLeuLysGluThrLeuGlu 2220
 Qy 2853 ACAAGCCCCACATCTGAGAGRTTCTCTAGT----- 2885
 Db 2221 LysThrGluIleAspPheTyrCysSerIleLeuIleSerGlnGlu 2240
 Qy 2886 ----- 2241 LysLeuGluLysAlaLysGluMetLeuGluThrGlnIleLeuLysGlu 2260
 Db 2919 CCAAGATAACCATATTCCATCACCAATGTCAGGAAACCGAAA 2918
 Db 2261 ----- 2269 ProLeuLeuGlyProVal---ValProGlyProSerProIleProSerValThrGluLys 2268
 Qy 2979 CCTACTCTCGCCGCGAAGGCCAAAGGCCAAAGTCAGGAG 3038
 Db 3039 AAGAGCCGGAAAGGGCGCTTGGGACAGG----- 3077

RESULT US-08-353-700-1
 Sequence 1, Application US/08353700
 Patent No. 5599919
 GENERAL INFORMATION:
 APPLICANT: YEN, TIMOTHY J.
 APPLICANT: RATTNER, JEROME B.
 TITLE OF INVENTION: NUCLEAR ACID ENCODING A
 TITLE OF INVENTION: TRANSIENTLY-EXPRESSED KINETOCHORE PROTEIN,
 TITLE OF INVENTION: AND METHODS OF USE
 NUMBER OF SEQUENCES: 4
 CORRESPONDENCE ADDRESS:
 ADDRESSEE: DANN, DORMAN, HERRELL AND SKILLMAN
 STREET: 1601 MARKET STREET, SUITE 720
 CITY: PHILADELPHIA
 STATE: PA
 COUNTRY: USA
 ZIP: 19103-2307

COMPUTER READABLE FORM:
 MEDIUM TYPE: Floppy disk
 COMPUTER: IBM PC compatible
 OPERATING SYSTEM: PC-DOS/MS-DOS
 SOFTWARE: PatentIn Release #1.0, Version #1.25
 CURRENT APPLICATION DATA:
 APPLICATION NUMBER: US/08/353,700
 FILING DATE: 09-DEC-1994
 CLASSIFICATION: 435
 ATTORNEY/AGENT INFORMATION:
 NAME: REED, JANET E.
 REGISTRATION NUMBER: 36,252
 TELECOMMUNICATION INFORMATION:

Db 2288 ArgLeuSerSerGlyGinAsnLysSerGlyLysSerGlyIleTrp 2307
 Qy 3078 -----CCTGCATCCCCATCCAATCATGCCGCTGCAACACT 3116
 Db 2308 GluAsnGlyGlyGlyProThrProAlaThrProGluSerPheSerLysLysLys 2327
 Qy 3117 GCAGCTCCACTGGAAATGCCGCTCTCTGATCTCAG----- 3155
 Db 2328 AlaValMetSerGlyIleHisProAlaGluAspThrGluLys 2347
 Qy 3156 -----GAAGTGCCTATGGGAGGACT----- 3176
 Db 2348 GlyLeuProGluValValLysGlyPheAlaAspIleProThrGlyLysThrSerPro 2367
 Qy 3177 -----ATCCTCAAGTCACCCGGAAACAACTGTTCCAGGCCAGGGAGTAC 3230
 Db 2368 TyrIleLeuArgGlyGluLysGlyPheAlaAspIleProThrGlyLys 2387
 Qy 3231 AACTCCATGCTAATCATCACCACGGACATAAAATTCACATTCACTGGGTCT 3290
 Db 2388 LeuAlaLeuSerProLeuSerSerLysGluIleLeuLys 2400
 .3291 CAGTTAACGGATCTGGGCTGCGCGTGAAGGCTGAGCCAGTTACCGTCCGG 3350
 Qy 3351 CCTGTCACGTCACGGAGGAGGTTCTACAGGCCAGTCCTCGCTCTCCAGG 3410
 Db 2415 LysValLysValAlaGlnArgSerProThrAlaGlyGlySerArgSer----- 2414
 Qy 3411 -----AACCACCTC-----TCTTCAGACCC 3431
 Db 2435 ThrLysSerValProValAsnLeuProGluArgSerProThrAspSerProArgGlu 2454
 Qy 3432 GGCTCTAGCAAGTGGACGAGCACTATAACCCGGTCACAACTGTCATCCACACGA 3491
 Db 2455 GlyLeuArgValLysBargGlyArgLeuValProSerProLysAlaGlyGlu 2474
 Qy 3492 GGAACCCAAATCAGTGTCAAGGACAA 3515
 Db 2475 GlySerGluAsnCysLysValGln 2482

TELEPHONE: (215) 563-4100
 TBLPAX: (215) 563-4044
 INFORMATION FOR SEQ ID NO: 1:
 SEQUENCE CHARACTERISTICS:
 LENGTH: 3248 amino acids
 TYPE: amino acid
 STRANDEDNESS: single
 TOPOLogy: linear
 MOLECULE TYPE: protein
 HYPOTHETICAL: NO
 ANTI-SENSE: NO
 ORGANISM: HUMAN
 i US-08-353-700-1

Alignment scores:

| Pred. No.: | Score: | Length: | Matches: | Conservative: | Mismatches: | Indels: | Gaps: |
|------------|----------|---------|----------|---------------|-------------|---------|-------|
| 1 | 1.17e-21 | 3248 | 316 | 240 | 527 | 450 | 63 |

US-10-788-793-1 (1-4364) x US-08-353-700-1 (1-3248)

Qy 51 TTAAGGAGTCGACAACAGGTGGGAATGAGATCAGGAAT-----CAAGGTGGAGAAAGT 104
 Db 1746 LeuSerSerArgSerLeuIeuLuglyileAspThrLeuAspAlaLeiGlnGlyArgAsnGlu 1765
 Qy 105 TCA-----TCTAACGGCATGTTCTCCCAACTCTCCATCATCAGCAGTGAT 155
 Db 1766 SerCysAspIleAspLysAspLysHisThrSerGluArgThrProLysAsp 1785
 Qy 156 GGTGCTAAAGGCCCTCAGAAGATGCAAAGAACACAAGGCCATCGAAAGGAGGGAT 215
 Db 1786 ValHisGlnIleCysAspIleAspLysAspAlaGlnGluAsp---LeuAsnLeuAspIle 1804
 Qy 216 GTCACTGGCTCCGAACTATCMAAGCACCTCAACCATTGGAGAA-----AGTGAG 269
 Db 1805 IleThrGluThrGlyAlaVal-----LysProThrGlyGluCysSerGlyGlu 1820
 Qy 270 AAAAGACTAACAGAGTCTGGAGTRATCCAAGGGAGCCTCATCCAGCTCTGAGTAC 329
 Db 1821 GlnSerProAspThrAsnTyroGluProProGlyGluAspLysThrGlySerSerGlu 1840
 Qy 330 ATGGIAGGGGAGTTGCAG-----GCT 350
 Db 1841 CysIleSerGluLeuSerPheSerAspIleProAsnAlaLeuValProMetAspHeLeuGly 1860
 Qy 351 CGAGAGATGTCCATCCAC-----ATGCTGAGGACAGAGAAAACCAAGGCCGAGGTCTG 404
 Db 1851 AsnGlnGluAspIleAsnLeuGlnSerAsnGluAsnIle 1880
 Qy 405 GAG---GCACACTATGGATCTGCAGAACCTGAGAAAGTCATGGCTCGGGTCTGACCGA--- 458
 Db 1881 ArgLeuLeuAspSerIleAsnValIleGluAspArgAspArgLysValGluSerLeuLeuAsn 1900
 Qy 459 -----GATGCC 464
 Db 1901 LysGluLeuAspSerIleAsnValIleGluAspArgLysValGluSerLeuLeuAsn 1920
 Qy 465 ATCCTTGCTCAAGAGAGAGTCATGGAGAACCTCTAGGAGCTATGAGAACTCTCAGAGCTG 524
 Db 1921 CysIleGluLeuGluIleAspSerIleValGlyGlu-----LeuIleLysGluAsnSerAspIle 1938
 Qy 525 GACAGACTGGAGAAAGCAGAAGGAGACGTACGCCGATCTCAGAGCTG 584
 Db 1939 SerGluIleAspLeuGluAspSerCysAspIleGluIleGluAspIleGluAspIle 1958
 Qy 585 GCTGAGAAGTGTACAGGCCACGGTGTACGGAGCTGGAGAAGCAGAACCCAG 644
 Db 1959 SerGlu-----GlyLeuAsnSerAspLeuGluMetIle 1970

Qy 645 GACTACATGAAACAAGAGCGGACACTTCACCAACCTG----- 680
 Db 1971 AspIleSerSerArgGluAspIleGlyAspAsnValAlaLysValAsnAspSerTrpIle 1990
 Qy 681 -----CTGGAGCAGGGAGAGAGCTG-----AAAAGCTCTT 716
 Db 1991 GluArgPheLeuAspValGluAsnGluLeuSerArgIleArgSerGluLysAlaSerIle 2010
 Qy 717 GAACAGAAAAGCTTACCAA----- 737
 Db 2011 GluIleGluAlaLeuTyroGluGluAlaAspIleGluValValGluThrGluLysLeuCys 2030
 Qy 738 GCCCGCAAAGAAACGCTAAGCCCT----- 770
 Db 2031 LeuGluIleAspAspAsnGluAsnLysGlnIleValIleValCysIleGluGluGluLeuSer 2050
 Qy 771 -----AACAACCTCGAATGAGCT-----GTGAAGCTCAAGTCC 806
 Db 2051 ValValThrSerGluArgAspGinLeuArgGlyGluLeuAspThrMetSerLysLysThr 2070
 Qy 807 TTGCCCCATGTTGGAGCAGGAGGGCAG----- 836
 Db 2071 ThrIleAlaIleAspGlnIleLeuAspIleValCysIleGluGluGluLeuSer 2090
 Qy 837 -----ATGCA----- 848
 Db 2091 GlnValThrSerGluArgAspGinLeuArgGlyGluLeuAspThrMetSerLysLysThr 2110
 Qy 849 CAACTGGAGCTGAGAGTCAGAAAGTCCAGGAGCTCTCAGAGCTGAGGGAGGGAA 908
 Db 2111 LeuIleGlnThrLeuSerSerAspValSerGluLeuLeuAspIle 2130
 Qy 909 GAAAGACTCAAGGGTCACTTACAATCCAAGGAGAACGCCAGAACGCTCAAGTTA 968
 Db 2131 GluIleAspIleGlnSerIle-----GluIleAspSerGlnAlaLeuSerLeuThr 2146
 Qy 969 GAAGTGGACTTGGAAACAAGGCCCTCGAGGTTCCAGGAGGCCAGAACGAGATGAACTGCC 1028
 Db 2147 LysCysGluIleAspGlnIleLeuAsnIleAspIleGlu----- 2162
 Qy 1029 AAATGGCGAATCAGAATCTCACAAACGGCAACTTCGACTCAA----- 1073
 Db 2163 ---IleLeuIleValGluSerGluSerLeuGlnAlaArgLeuSerAspPheGlu 2181
 Qy 1074 -----CTGGIT 1079
 Db 2182 LysLeuAsnValSerIleAsnAlaLeuGluAlaAlaLeuValGluIleGlyGluPheAlaIle 2201
 Qy 1080 GGCTTATCGCAAAGGATGGAGGAGACCAATAAGCCTCAGAG----- 1133
 Db 2202 ArgLeuSerSerThrGlyGluIleGluValGlnLeuAspSerIleLeuAsn 2221
 Qy 1134 -----GCAAGGAGAGCTCCAG-----GAGCTGAGAGAAATGCCAAGGG 1178
 Db 2222 ArgLeuSerSerThrGlyGluIleGluValGlnLeuAspSerIleLeuAsn 2241
 Qy 1179 GAATGCGAACTCCAGTCATGGCGGAGTGGAGAGCTGCGCAAGGCC 1229
 Db 2242 GluArgGluAsnAspSerIleGluValGlnLeuAspSerIleLeuAsn 2281
 Qy 1230 -----GTCCTGAGATGGAGGAGGAGCTCAAGAGAACCCAG 1265
 Db 2262 SerGluGluAsnGlnIleGluLeuValIleLeuAspAlaGluAsnSerIleGlu 2281
 Qy 1266 AAGACGGGCCAGTGGCGGAGCTGAGAGAACGCTCCAAGAGGAAGAACCCAGC 1325
 Db 2282 ThrLeuIleSerGlyGluIleLeuAsnSerAspIleGluAspIle 2301
 Qy 1326 AAGGAACCTAGACTAGAAGTGGAGAAGCTGCGAGAG-----AGG 1364
 Db 2302 ValThrIleArgSerGlyGluIleAsnLeuThrIleGlyGluIleGlyGlu 2321
 Qy 1365 ATGCTGAGCTGGAGAGCTGGAGGAAGCGTTCAAGCCGAGTAGTCGGAAATGCCACCCAG 1424

| | | |
|----|------------------------------------------------------------------------|------|
| Db | 2322 LeuSerGluLeuAspLysLeuSerSerPheLysSerLeuLeuGluLysGluIn | 2341 |
| Qy | 1425 CTCATCTGAACTGGAGAGAGAACCTAACAGACCTGCTG--AACGAGCTG | 1481 |
| Db | 2342 AlaGluIleGlnIleLysGluGluSerLysThrAlaValGluMetLeuGlnAsglnIeu | 2361 |
| Qy | 1482 GAGGTGGTCAAGACTCGGAGTAAAGAACTCGATGCTCCAGAGGTAGACTCGAGAGGCC | 1541 |
| Db | 2362 LysGluLeuAspGluValAlaAlaLeu--CygLyAspGlnGluMetLysAla | 2380 |
| Qy | 1542 --ArgLeuGluAspProProLeuGluGluLysGlnLeuArgAsnSerIleGlu | 2400 |
| Db | 2381 ThrGluGlnSerLeuAspProProLeuGluGluLysGlnLeuArgAsnSerIleGlu | 2400 |
| Qy | 1569 AAGCTGAAGTCCTCACTGTGATGCTGGATGGAGAGGAAATATG----ATGGAG | 1622 |
| Db | 2401 LysLeuArgAla---ArgLeuGluAlaAspGluLysGlnLeuCysValLeuGln | 2418 |
| Qy | 1623 AAATAAAGCAAGAGAGAGAACTGGATGGTTGAATAAACTTAAGGTGGAGCAG | 1682 |
| Db | 2419 GlnLeuLysGluSerGluLysAlaAspGluLysGlnLeuGlnAsglnIleGlu | 2431 |
| Qy | 1683 GAAAGTCACTGGATGTGACGGAAAGLACTGAA- 1712 | 1712 |
| Db | 2432 GlyArgValGluAsnLeuGluArgGluLeuGluLysGlnLeuAsnSerGluLys | 2431 |
| Qy | 1713 -----ATCGAGGAAGCAAG-----AAGCTTTAACCTCAATCTGAA-- 1751 | 1751 |
| Db | 2452 AlaleuGluAlaGluAsnSerLysGlyGluValGluThrLeuLysAlaLysIleGlu | 2471 |
| Qy | 1752 -----ATGGAGGAAGAGGACTACAGTCTGACAAGAGGGAT 1790 | 1790 |
| Db | 2472 MetThrGlnSerLeuArgGlyLeuAspValValThrIleArgSerLysGlu | 2491 |
| Qy | 1791 GAGCTGATGGTAAGTGAGGAGCTAGAGCTTCCAAGGACTACAGCCAGCTCAGGAG 2435 | 2435 |
| Db | 2492 AsnLeuAspCysMetHisLysAspGlnValGluLysGluIleValArgGlu | 2791 |
| Qy | 2792 IleAlaGluTyrglnLeuArgGluLysGluAlaGluLysGluIleValLeuIleGlu | 2791 |
| Db | 2792 IleAlaGluTyrglnLeuArgGluLysGluAlaGluLysGluIleValLeuIleGlu | 2811 |
| Qy | 2436 AGTGGAAACGGCCGAAGGATGTTGACCTGCTCCACTGGGAGCTCAGGGAG 2495 | 2495 |
| Db | 2812 AspThrAsnLysGlnTyrgluValGluLysGluAsnSerGluLys | 2831 |
| Qy | 2496 GCGGTGTGGGGATGCTGGAGGGAGACCCGGCTGTCATCGCAATCRTC 2555 | 2555 |
| Db | 2832 GluGluCysAsnSerSerGlnLysGlu-----ileAspLeuLysSer | 2848 |
| Qy | 2556 CAGGAGAA-----AATCACATC----- 2573 | 2573 |
| Db | 2849 LysGluGluLeuAsnSerSerLeuLysGluLysGlu 2868 | 2868 |
| Qy | 2512 SerSerPheGluAsnIleLeuGluLysGluGlnGluAsnSerGlnLeuGluLys | 2511 |
| Db | 1838 ----- 1838 | 1838 |
| Qy | 2574 -----ATGAGTAATCTCGACAGTAGGC--CTGAGAACCCATGGAACGGTC-- 2621 | 2621 |
| Db | 2532 SerSerThrAlaMetGluMetLeuGlnThrGlnIleLysGluAsnGlnMetLysGlu | 2531 |
| Qy | 1839 --AGCTGCACTGAGACTACTAAAGAGCCGCTGATGGCATA--GAGGAGTAGAA 1892 | 1892 |
| Db | 2669 ThrLysMetAspAsnLeuAsnSerSerLeuLysAlaThrThrIleLeuGluLysGlu | 2888 |
| Qy | 2552 AlaLeuIleAsnAspGlnGluAlaCysLysAlaLysGluGlnAsnLeuAsnSerGlnVal 2571 | 2571 |
| Db | 2572 GluCysLeuGluGluGluLysGluGlnAsnLeuAsnSerGlnVal 2571 | 2571 |
| Qy | 1935 ACCTGCCCCGGAA 1946 | 1946 |
| Db | 2572 GluCysLeuGluGluGluLysGluGlnAsnLeuAsnSerGlnVal 2571 | 2571 |
| Qy | 2592 TyrIleValLeuGluGlnGluValGluAspGlyLysGln 2611 | 2611 |
| Db | 2592 TyrIleValLeuGlnSerValLysGlyLeuIleGlnGlu 2611 | 2611 |
| Qy | 1953 AAGATCAGAGAACTAACCGCTGAAATCGAGAGACTGAGAACGGCTCCAGCTGGAG 2012 | 2012 |
| Db | 2612 LysLeuGluLysLysAspGluGluIleSerArgLeuLysAsnGlnAspGlnIeu | 2631 |
| Qy | 2013 GTGGTGGAGGGGACTGTGAGAGACCGAGGATAAG-- 2054 | 2054 |
| Db | 2632 GluLeuValSerLysLeuSerGlnValGluLysGlnAsn 2651 | 2651 |
| Qy | 2055 -----CAGTGGAGGAGGTTCAAGACCGAGGATAAG-- 2090 | 2090 |
| Db | 2652 LeuGluLeuGluAsnLeuLeuGluGluGluLysGlu 2671 | 2671 |
| Qy | 2091 GCAAACCTCTCCCAGCAGCTGAGGAATCAAACAC----- 2129 | 2129 |

ISBN-0-7887-9311-7

Page 12

TELEFAX: (215) 563-4100
INFORMATION FOR SEQ ID NO: 1:
SEQUENCE CHARACTERISTICS:
LENGTH: 3248 amino acids
TYPE: amino acid
STRANDEDNESS: not relevant
TOPOLOGY: not relevant
MOLECULE TYPE: protein
HYPOTHETICAL: NO
ARTICLE: NO

| US-10-788-793-1 (1-4364) x PCT-US95-16216-1 (1-3248) | | Length: | 3248 |
|------------------------------------------------------------------------|------------------------------------------------------------------------|---------------|------|
| Qy | Db | Matches: | 316 |
| Qy | Db | Conservative: | 240 |
| Qy | Db | Mismatches: | 527 |
| Qy | Db | Indels: | 450 |
| Qy | Db | Gaps: | 63 |
| 51 TTAAGGAGTCGACAACAGGTGGGAATCGAGATCACCAAAT-----CAAGGTGGAGAAAGT 104 | 1746 IeuserSerargSerLeuLeuGlyIleAspThrGluAspAlaLeuGlyArgAsnGlu 1765 | | |
| 1786 ValHisGinileCysAspLysAspAlaIleGlnAsp---LeuAsnLeuAspIleGluLys 1785 | 216 GTCATGGCTTCGGAACATCAAAGGCACCTCAACCATCTGGAGAA----AGTAG 215 | | |
| 1805 IleThrGluThrGlyIlaVal-----LysProThrGlyGlu 269 | 1805 IleThrGluThrGlyIlaVal-----LysProThrGlyGlu 269 | | |
| 270 AAAAGACTAACGAAACTCTGTGGAGTTATCCAAGGAGCTCATCCAGCTCTGAGTAC 329 | 1821 GlnSerProAspThrAsnTyrgluProProGlyGlu 1820 | | |
| 330 ATGGAAGGGAGGTGCAAG-----GCT 350 | 330 ATGGAAGGGAGGTGCAAG-----GCT 350 | | |
| 1841 CysIleSerGluLeuSerPheSerGlyProAsnAlaLeuValProMetAspPheLeuGly 1860 | 351 CGAGAACATGTCATCCAC-----ATGCTGAGGACAGAGAAACCAAGGCCAGGTCTG 404 | | |
| 1861 AsnGlnGluAspIleHisAsnLeuGlnLeuArgValLysGluThrSerAsnGluAsnLeu 1880 | 405 GAG---GCACACTATGGATCTGAGAACCTGAGAAAGTGCTTCGGTCTGACCGA--- 458 | | |
| 459 ----- | 1881 ArgLeuLeuHisValIleGluAspArgAspArgLysValGluSerLeuLeuAsnGluMet 1900 | | |
| 1901 LysGluLeuAspSerLysLeuHisLeuGlnGluValGlnLeuMetThrLysIleGluIla 1920 | 465 ATCCTTGCTCAAGAGAACGCTACGGAGAGACGGCTATGAGAACCTATCTGAGAGCTG 524 | | |
| 1921 CysIleGluLeuGluLysIleValGlyGlu-----LeuLysGluAsnSerAspLeu 1938 | 525 GACAGACTGGAAAGCAGAACGAGACCTACGGGCGATGAGGAGCTGGAGAAGCACAAGCAC 584 | | |
| 1939 SerGluLysLeuGluThrPheSerCysAspHisGlnGluLeuGlnArgValGluThr 1956 | 585 GCTGAGGAAGTGTCAAGGGCACCGTGTACGAGCTGGAGAAGCACAAGCAC 644 | | |
| 1959 serGlu-----GluGlu-----GluGlu-----GluGlu-----GluGlu-----Glu 63 | 1959 serGlu-----GluGlu-----GluGlu-----GluGlu-----GluGlu-----Glu 63 | | |